PA NT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference		of Transmittal of International Search Report 220) as well as, where applicable, item 5 below.
03-004-B	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
International application No.	International filing date (day/month/year)	(Lamesty) Horry Bato (Bay/Horralysal)
PCT/US 03/36747	13/11/2003	12/11/2002
Applicant		
RIGEL PHARMACEUTICALS, IN	<u>c. </u>	
This International Search Report has bee according to Article 18. A copy is being tr	n prepared by this International Searching Au ansmitted to the International Bureau.	thority and is transmitted to the applicant
This International Search Report consists It is also accompanied by	s of a total of8 sheets. • a copy of each prior art document cited in thi	s report.
Basis of the report		•
a. With regard to the language, the language in which it was filed, un	international search was carried out on the balless otherwise indicated under this item.	asis of the international application in the
the international search v Authority (Rule 23.1(b)).	vas carried out on the basis of a translation of	the international application furnished to this
b. With regard to any nucleotide ar was carried out on the basis of the	nd/or amino acid sequence disclosed in the sequence listing:	international application, the international search
contained in the internation	onal application in written form.	
filed together with the into	ernational application in computer readable for	rm.
furnished subsequently to	o this Authority in written form.	
furnished subsequently to	o this Authority in computer readble form.	
	bsequently furnished written sequence listing as filed has been furnished.	does not go beyond the disclosure in the
the statement that the inf furnished	ormation recorded in computer readable form	is identical to the written sequence listing has been
2. Certain claims were fou	und unsearchable (See Box I).	
3. Unity of invention is lac	king (see Box II).	
4. With regard to the title ,		
the text is approved as s	ubmitted by the applicant.	
X the text has been establi	shed by this Authority to read as follows:	
RHODANINE DERIVATIVES	AND PHARMACEUTICAL COMPOSI	TIONS CONTAINING THEM
5. With regard to the abstract,	·	
	ubmitted by the applicant.	
the text has been establi within one month from the	shed, according to Rule 38.2(b), by this Authore date of mailing of this international search re	ority as it appears in Box III. The applicant may, eport, submit comments to this Authority.
6. The figure of the drawings to be put	olished with the abstract is Figure No.	
as suggested by the app	licant.	None of the figures.
because the applicant fa	iled to suggest a figure.	
because this figure bette	er characterizes the invention.	

Inte

ional application No.

INTERNATIONAL SEARCH REPORT

PCT/US 03/36747

Box III TEXT OF THE ABSTRACT (Continuation of item 5 of the first sheet)

This invention describes rhodanine derivatives and pharmaceutical compositions useful as inhibitors of ubiquitination. The compounds and compositions of the invention are useful as inhibitors of the biochemical pathways of organisms in which ubiquitination is involved. In particular, the compounds and compositions are useful for treating cell proliferative diseases such as cancers.

Internat 1 Application No PCT/US 03/36747

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C07D417/06 C07D417/14 A61K31/427 A61P35/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C07D

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, CHEM ABS Data

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 02 051409 A (KYOWA HAKKO KOGYO KK; AKAMA TSUTOMU (US); GERON CORP (US); HOLCOMB) 4 July 2002 (2002-07-04) examples 1-5, 7, 10-13, 17, 18, 28, 30, 32, 34-36, 39, 40, 43-47, 54-56 claims 1,22-30	1-5,11, 15-17
X	WO 02 053155 A (YAMASHITA YOSHINORI; HAGIHARA KOJI (JP); ARAI HITOSHI (JP); ASAI A) 11 July 2002 (2002-07-11) page 26 -page 75; claims	1-11, 15-17
X	WO 98 53790 A (BIEDIGER RONALD J; SCOTT IAN L (US); MARKET ROBERT V (US); TEXAS B) 3 December 1998 (1998-12-03) page 10, line 16 - line 20; claims	1-11, 15-17

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents: A* document defining the general state of the art which is not considered to be of particular relevance E* earlier document but published on or after the international filing date L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) O* document referring to an oral disclosure, use, exhibition or other means P* document published prior to the international filing date but later than the priority date claimed	 'T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention 'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone 'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combined with one or more other such documents, such combination being obvious to a person skilled in the art. '&' document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
22 March 2004	29/03/2004
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk	Authorized officer
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Hass, C

Interna' I Application No PCT/ປຣ 03/36747

C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 00 10573 A (BAILEY THOMAS R; VIROPHARMA INC (US); YOUNG DOROTHY C (US)) 2 March 2000 (2000-03-02) claims 1,17,19,21,22,29,30,36,37,47; tables I,II,IV,V,VI	1-11
X	WO 00 32598 A (STRUCTURAL BIOINFORMATICS INC) 8 June 2000 (2000-06-08) figure pages 1/25 to 23/25	1-11
X	WO 01 77091 A (JAEN JUAN C; LI YANG (US); PIPER DEREK E (US); TULARIK INC (US); P) 18 October 2001 (2001-10-18) figure pages 1/20 to 19/20 claims 23-25	1-4,11
X	EP 0 587 377 A (LILLY CO ELI) 16 March 1994 (1994-03-16) examples 2, 3, 6, 9, 10, 13-16, 22, 23, 33-36, 39, 75, 76	1-4,11
X	EP 1 207 155 A (SHIONOGI & CO) 22 May 2002 (2002-05-22) page 39, compound no. 3	1,2,11
X	US 2 824 087 A (UNRUH CORNELIUS C ET AL) 18 February 1958 (1958-02-18) table I, examples 4-6, 9, 11	1,2,11
X	EP 0 976 748 A (SHIONOGI & CO) 2 February 2000 (2000-02-02) page 40, reaction 8 -> 9 and page 42, lines 35, 36, compound 4-(2,4-dioxothiazolidine-5-ylidenemethyl)- benzoic acid	1,2,11
X	US 5 554 767 A (YUEN PO-WAI ET AL) 10 September 1996 (1996-09-10) examples 1, 2, 7, 9, 11, 15, 19, 21, 24, 26, 28, 30, 35, 37, 39, 41, 48, 50, 52	1,2,11
X	EP 0 343 643 A (WARNER LAMBERT CO) 29 November 1989 (1989-11-29) examples 7, 8, 17, 26-32, 34, 38-41, 43, 54, 57, 61, 65, 104, 105	1,2,11
X	WO 00 18748 A (SCHAEFER WOLFGANG; HOFFMANN EIKE (DE); HONOLD KONRAD (DE); KALUZA) 6 April 2000 (2000-04-06) pages 10 to 16, compounds 1-4, 21, 22, 33, 41, 44, 46, 49, 54-57, 59, 60, 65, 74, 82, 84, 97, 100, 103, 106, 110, 114, 116, 134-136, 170, 181, 186, 195, 196, 198-200, 202, 209, 210, 215, 216	1,2,11
	-/	

2

Interna' I Application No
PCT/US 03/36747

lation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
WO 00 18746 A (SCHAEFER WOLFGANG; HONOLD KONRAD (DE); KALUZA KLAUS (DE); ESSWEIN) 6 April 2000 (2000-04-06) pages 9 to 16, compounds 1-3, 5, 6, 27, 28, 42, 56, 58, 62, 70-72, 74-77, 81, 99, 101, 107, 114, 117, 119, 120, 125, 129, 130, 132, 137, 151, 157-159, 188, 198, 214, 220, 229, 231, 237, 238	1,2,11
US 4 358 521 A (KONDO SYUNICHI ET AL) 9 November 1982 (1982-11-09) column 11, example 11	1,2
US 3 678 041 A (MOUSSERON MAX J) 18 July 1972 (1972-07-18) tables I,II,III	1,2
WO 01 10881 A (3M INNOVATIVE PROPERTIES CO) 15 February 2001 (2001-02-15) page 52, Sequence 3 page 52, line 5 - line 10	1-4
GB 2 387 172 A (PANTHERIX LTD) 8 October 2003 (2003-10-08) table 1	1-11
GB 2 386 892 A (PANTHERIX LTD) 1 October 2003 (2003-10-01) page 7	1-11
WO 03 072033 A (PELLECCHIA MAURIZIO; CHANG EDCON (US); DONG QING (US); HANSEN MARK) 4 September 2003 (2003-09-04) claims 1,2,5-11,20,21,23,25-27,34,45,46,49,51,52	1-10
WO 03 027081 A (KAARSHOLM NIELS C; JAKOBSEN PALLE (DK); LUDVIGSEN SVEND (DK); MADS) 3 April 2003 (2003-04-03) examples 60, 61, 63, 64, 68, 70, 74, 76, 78, 81, 83, 87, 91-96, 98, 110, 114-117, 124, 125, 130-132, 135, 136, 141, 144, 148, 151, 154, 155, 159-162, 164-166, 170, 173, 186, 192, 206, 207, 213, 216, 224, 227, 229, 233, 234, 236, 260, 262, 263, 269, 274	1,2,11
WO 03 043998 A (MAXIA PHARMACEUTICALS INC) 30 May 2003 (2003-05-30) claims 1,42-48; examples	1-11, 15-17
	WO 00 18746 A (SCHAEFER WOLFGANG; HONOLD KONRAD (DE); KALUZA KLAUS (DE); ESSWEIN) 6 April 2000 (2000-04-06) pages 9 to 16, compounds 1-3, 5, 6, 27, 28, 42, 56, 58, 62, 70-72, 74-77, 81, 99, 101, 107, 114, 117, 119, 120, 125, 129, 130, 132, 137, 151, 157-159, 188, 198, 214, 220, 229, 231, 237, 238 US 4 358 521 A (KONDO SYUNICHI ET AL) 9 November 1982 (1982-11-09) column 11, example 11 US 3 678 041 A (MOUSSERON MAX J) 18 July 1972 (1972-07-18) tables I,II,III WO 01 10881 A (3M INNOVATIVE PROPERTIES CO) 15 February 2001 (2001-02-15) page 52, Sequence 3 page 52, line 5 - line 10 GB 2 387 172 A (PANTHERIX LTD) 8 October 2003 (2003-10-08) table 1 GB 2 386 892 A (PANTHERIX LTD) 1 October 2003 (2003-10-01) page 7 WO 03 072033 A (PELLECCHIA MAURIZIO; CHANG EDCON (US); DONG QING (US); HANSEN MARK) 4 September 2003 (2003-09-04) claims 1,2,5-11,20,21,23,25-27,34,45,46,49,51,52 WO 03 027081 A (KAARSHOLM NIELS C; JAKOBSEN PALLE (DK); LUDVIGSEN SVEND (DK); MADS) 3 April 2003 (2003-04-03) examples 60, 61, 63, 64, 68, 70, 74, 76, 78, 81, 83, 87, 91-96, 98, 110, 114-117, 124, 125, 130-132, 135, 136, 141, 144, 148, 151, 154, 155, 159-162, 164-166, 170, 173, 186, 192, 206, 207, 213, 216, 224, 227, 229, 233, 234, 236, 260, 262, 263, 269, 274 WO 03 043998 A (MAXIA PHARMACEUTICALS INC) 30 May 2003 (2003-05-30)

2

ional application No. PCT/US 03/36747

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. χ	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Although claims 12-17 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	on Protest The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.

Informati. n patent family members

Interna 1 Application No PCT/US 03/36747

			·-·		'	01/03	03/30/4/
	atent document d in search report		Publication date		Patent family member(s)		Publication date
WO	02051409	A	04-07-2002	US WO	2002120144 02051409		29-08-2002 04-07-2002
WO	02053155	Α	11-07-2002	МÓ	02053155	A1	11-07-2002
WO	9853790	Α	03-12-1998	AU WO	7684798 9853790		30-12-1998 03-12-1998
WO	0010573	Α	02-03-2000	AU AU	743411 5570299		24-01-2002 14-03-2000
				BR	9913157	Α	15-05-2001
				CA	2341970		02-03-2000
				CN	1325309		05-12-2001
				EP	1128832		05-09-2001
				ID	27787		26-04-2001
				JP	2002523371		30-07-2002
				WO	0010573		02-03-2000
				US	2003195213		16-10-2003
WO	0032598	A	08-06-2000	AU	3111700	Α	19-06-2000
		· 		WO	0032598		08-06-2000
WO	0177091	Α	18-10-2001	AU	5320601	Α	23-10-2001
,		. •		WO	0177091		18-10-2001
				US	2002142290		03-10-2002
EP	0587377	 -	16-03-1994	AU	676843		27-03-1997
				ΑU	4621893		17-03-1994
				CA	2105598		11-03-1994
				CN	1091006		24-08-1994
				CZ	9301814		16-03-1994
				EP	0587377		16-03-1994
				EP	0915090		12-05-1999
				FI HU	933946 70184		11-03-1994
	•			IL	106877		28-09-1995 10-03-1998
				IL	119119		16-08-1998
				JP	6192091		12-07-1994
				MX	9305444		31-05-1994
				NO	933198		11-03-1994
				NO	981911		11-03-1994
				NZ	248573		27-02-1996
				PL	300335		21-03-1994
				RU	2131251		10-06-1999
				US	5523314		04-06-1996
				US	5716975		10-02-1998
				US	5661168		26-08-1997
				ZA 	9306492 	A 	02-03-1995
EP	1207155	Ą	22-05-2002	AU	6023300		13-02-2001
				CA	2380206		01-02-2001
				EP	1207155		22-05-2002
				CN	1376150		23-10-2002
				WO	0107423	Al 	01-02-2001
US	2824087 <i>F</i>	A	18-02-1958	NONE			
EP	0976748 <i>F</i>	A	02-02-2000	AT	255579	T	15-12-2003

Informati a patent family members

Interna' N Application No PCT/しら 03/36747

Patent document					03/36/4/
cited in search report	Į	Publication date		Patent family member(s)	Publication date
EP 0976748	A		AU	719210 B2	04-05-2000
			AU	5577598 A	25-08-1998
			BR	9807132 A	25-01-2000
	•		CA	2277947 A1	06-08-1998
			DE	69820248 D1	15-01-2004
			EP	0976748 A1	
					02-02-2000
			HU	0000965 A2	28-04-2001
			NO	993706 A	30-09-1999
			US	6147100 A	14-11-2000
			CN	1251587 T	26-04-2000
			WO	9833797 A1	06-08-1998
			PL	334915 A1	27-03-2000
			RU	2198174 C2	10-02-2003
			TR	9901847 T2	21-06-2000
US 5554767	A	10-09-1996	US	5760048 A	02-06-1998
EP 0343643	Α	29-11-1989	ΑT	103175 T	15-04-1994
		_: _=	ΑU	626863 B2	13-08-1992
			AU	3505889 A	30-11-1989
			CA	1340247 C	15-12-1998
			DE	68914029 D1	28-04-1994
			DE	68914029 T2	07-07-1994
			DK		
				252089 A	26-11-1989
			EP	0343643 A2	29-11-1989
			EP	0565135 A1	13-10-1993
			ES	2063073 T3	01-01-1995
			FΙ	892522 A	26-11-1989
			ΙE	62214 B1	11-01-1995
			ΙE	940525 L	25-11-1989
			JP	2062864 A	02-03-1990
			JP	2899309 B2	02-06-1999
			KR	9702228 B1	26-02-1997
			NO	892083 A	27-11-1989
			NZ	229266 A	23-12-1991
			PH	27092 A	26-02-1993
			1 11	Z/U3/ A	
			PT	90662 A ,B	30-11-1989
			PT US	90662 A ,B 5464856 A	30-11-1989 07-11-1995
			PT US US	90662 A ,B 5464856 A 5208250 A	30-11-1989 07-11-1995 04-05-1993
			PT US US US	90662 A ,B 5464856 A 5208250 A 5306822 A	30-11-1989 07-11-1995 04-05-1993 26-04-1994
			PT US US US ZA	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
 WO 0018748	A	06-04-2000	PT US US US ZA AU	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
			PT US US US ZA AU WO	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018748	A A	06-04-2000	PT US US US ZA AU WO	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
			PT US US US ZA AU WO	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
			PT US US ZA AU WO AU WO	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746	A	06-04-2000	PT US US ZA AU WO AU WO	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746	A	06-04-2000	PT US US US ZA AU WO AU WO JP DE	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746 US 4358521	A A	06-04-2000	PT US US ZA AU WO AU WO JP DE FR BE	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1 2081553 A6 716140 A	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746 US 4358521	A A	06-04-2000	PT US US US ZA AU WO AU WO JP DE	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746 US 4358521	A A	06-04-2000	PT US US ZA AU WO AU WO JP DE FR BE	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1 2081553 A6 716140 A	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746 US 4358521	A A	06-04-2000	PT US US ZA AU WO AU WO JP DE FR BE DE	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1 2081553 A6 716140 A 1770583 A1	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991
WO 0018746 US 4358521	A A	06-04-2000	PT US US US ZA AU WO AU WO JP DE FR BE DE FR	90662 A ,B 5464856 A 5208250 A 5306822 A 8903834 A 6330999 A 0018748 A1 6331099 A 0018746 A1 56156832 A 3117701 A1 2081553 A6 716140 A 1770583 A1 6797 M	30-11-1989 07-11-1995 04-05-1993 26-04-1994 30-01-1991

 Interna 1 Application No
PCT/US 03/36747

	atent document I in search report		Publication date		Patent family member(s)		Publication date
US	3678041	A		US ES	3704296 354732		28-11-1972 16-04-1970
WO	0110881	A	15-02-2001	US AU BR DE EP JP WO US	6372895 6614100 0012937 60006970 1204671 2003506455 0110881 2002147317	A A D1 A1 T A1	16-04-2002 05-03-2001 30-04-2002 15-01-2004 15-05-2002 18-02-2003 15-02-2001 10-10-2002
GB	2387172	Α	08-10-2003	NONE			
GB	2386892	Α	01-10-2003	NONE			
WO	03072033	Α	04-09-2003	US WO	2004009526 03072033		15-01-2004 04-09-2003
WO	03027081	Α	03-04-2003	WO US	03027081 2003229120		03-04-2003 11-12-2003
WO	03043998	A	30-05-2003	WO US WO US WO US	03043998 2003144329 03075924 2003216432 03075858 2004034004	A1 A1 A1 A2	30-05-2003 31-07-2003 18-09-2003 20-11-2003 18-09-2003 19-02-2004